WE CLAIM:

1. A compound of general formula:

Wherein:

 R_1 is selected from H and CH_3 , and R_2 is selected from H and OH, or R_1 and R_2 together form an optionally substituted phenyl ring which is fused to the pyridine ring; and

R₃ is selected from H, CH₃, CH₂OH and

R₄ is selected from H, CH₃, CH₂OH,

$$X$$
 Y
 O
 OR_6
 OR_7
 OR_7

R₅ is selected from H, phenyl, halogen-substituted phenyl and

Wherein R_6 and R_7 are each independently selected from H, Na^+ , K^+ , alkyl and optionally substituted aryl, and X and Y are each independently selected from H, OH and F, or at least one of X and Y is an heteroatom and together with R_3 forms a bridge with the proviso that R_4 is

$$X$$
 Y O OR_6 OR_7 OR_7

and N-oxides thereof, and biologically acceptable salts thereof.

- 2. The compound according to claim 1, wherein said halogen-substituted phenyl is a fluoro-substituted phenyl.
- 3. The compound according to claim 1, wherein said halogen-substituted phenyl is $p-C_6H_4F$.
- 4. The compound according to any one of claims 1 to 3, wherein said heteroatom is selected from O and S.

- 5. The compound according to any one of claims 1 to 3, wherein said heteroatom is O.
- 6. The compound according to any one of claims 1 to 5, wherein said bridge is selected from $-CH_2-$, $-CH_2CH_2-$ and $-CH_2CH_2-$.
- 7. The compound according to any one of claims 1 to 5, wherein said bridge is a methylene bridge.
- 8. The compound according to any one of claims 1 to 7, wherein said alkyl is a C_1 to C_6 straight or branched alkyl.
- 9. The compound according to any one of claims 1 to 7, wherein said alkyl is t-butyl.
- 10. The compound according to any one of claims 1 to 9, wherein said aryl is phenyl or naphthyl.
- 11. The compound according to claim 1, wherein R_1 , R_2 , R_4 , R_5 are all H and R_3 is

12. The compound according to claim 1, wherein R_1 , R_2 , R_3 , R_5 are all H and R_4 is

13. The compound according to claim 1, wherein R_1 , R_2 , R_3 , R_4 are all H and R_5 is

14. The compound according to claim 1, wherein R_1 and R_2 together form an optionally substituted phenyl ring which is fused to the pyridine ring; R_3 and R_5 are both H; and R_4 is

15. The compound according to claim 1, wherein R_1 and R_3 are both CH_3 ; R_2 is OH; R_5 is H; and R_4 is

16. The compound according to claim 1, wherein R_1 and R_4 are both CH_3 ; R_2 is OH; R_5 is H; and R_3 is

17. The compound according to claim 1, wherein R_1 is CH_3 ; R_2 is OH; R_3 is CH_2OH ; R_5 is H; and R_4 is

18. The compound according to claim 1, wherein R_1 is CH_3 ; R_2 is OH; R_3 is CH_2OH ; R_5 is H; and R_4 is

19. The compound according to claim 1, wherein R_1 is CH_3 ; R_2 is OH; R_3 is CH_2OH ; R_5 is C_6H_5 ; and R_4 is

20. The compound of claim 1, wherein R_1 is CH_3 ; R_2 is OH; R_3 is CH_2OH ; R_5 is $p-C_6H_4F$; and R_4 is

- 21. A compound according to claim 1, wherein R₅ is p-C₆H₄F.
- 22. A compound according to claim 1 selected from: [Hydroxy-(5-hydroxy-4hydroxymethyl-6-methyl-2-phenyl-pyridin-3-yl)-methyl]-phosphonic acid; {[2-(4-Fluoro-phenyl)-5-hydroxy-4-hydroxymethyl-6-methyl-pyridin-3-yl]-hydroxymethyl}phosphonic acid; [Hydroxy-(4-pyridin-2-yl-phenyl)-methyl]-phosphonic acid; [Fluoro-(4-pyridin-2-yl-phenyl)-methyl]-phosphonic acid; (Hydroxy-quinolin-3-yl-methyl)phosphonic acid; (Fluoro-quinolin-3-yl-methyl)-phosphonic acid; [Hydroxy-(5hydroxy-4,6-dimethyl-pyridin-3-yl)-methyl]-phosphonic acid; (Hydroxy-pyridin-4-ylmethyl)-phosphonic acid; (Hydroxy-pyridin-3-yl-methyl)-phosphonic acid; (3,7-Dihydroxy-6-methyl-1,3-dihydro-furo[3,4-c]pyridin-3-yl)-phosphonic acid: [(3,7-Dihydroxy-6-methyl-1,3-dihyrdo-furo[3,4-c]pyridin-3-yl)-difluoromethyl]-phosphonic acid; and nicotinyl phosphonates thereof, N-oxides thereof, phosphonate esters thereof and biologically acceptable salts thereof.

23. A compound according to claim 1 comprising:

- 24. A pharmaceutical composition comprising a therapeutically effective amount of a compound according to any one of claims 1 to 23 and a pharmaceutically acceptable carrier.
- 25. The compound according to any one of claims 1 to 24, wherein at least one polar group is blocked by a lipophilic moiety capable of being enzymatically cleaved off after absorption into the circulatory system.
- 26. The compound according to claim 25, wherein said lipophilic moiety is an ester.
- 27. The compound according to claim 25, wherein said lipophilic moiety is a phosphonate ester.
- 28. A method of treating hypertension in a mammal comprising administering to the mammal a therapeutically effective amount of a compound according to any one of claims 1 to 27 in a unit dosage form.

- 29. A method of treating myocardial infraction in a mammal comprising administering to the mammal a therapeutically effective amount of a compound according to any one of claims 1 to 27 in a unit dosage form.
- 30. A method of treating ischemia reperfusion injury in a mammal comprising administering to the mammal a therapeutically effective amount of a compound according to any one of claims 1 to 27 in a unit dosage form.
- 31. A method of treating myocardial ischemia in a mammal comprising administering to the mammal a therapeutically effective amount of a compound according to any one of claims 1 to 27 in a unit dosage form.
- 32. A method of treating congestive heart failure in a mammal comprising administering to the mammal a therapeutically effective amount of a compound according to any one of claims 1 to 27 in a unit dosage form.
- 33. A method of treating arrhythmia in a mammal comprising administering to the mammal a therapeutically effective amount of a compound according to any one of claims 1 to 27 in a unit dosage form.
- 34. A method of reducing blood clots in a mammal comprising administering to the mammal a therapeutically effective amount of a compound according to any one of claims 1 to 27 in a unit dosage form.
- 35. A method of treating hypertrophy in a mammal comprising administering to the mammal a therapeutically effective amount of a compound according to any one of claims 1 to 27 in a unit dosage form.
- 36. A method of treating a disease that arises from thrombotic and prothrombotic states in which the coagulation cascade is activated in a mammal comprising

administering to the mammal a therapeutically effective amount of a compound according to any one of claims 1 to 27 in a unit dosage form.

- 37. A method of treating diabetes mellitus in a mammal comprising administering to the mammal a therapeutically effective amount of a compound according to any one of claims 1 to 27 in a unit dosage form.
- 38. A method of treating insulin resistance in a mammal comprising concurrently administering to the mammal a therapeutically effective amount of a compound according to any one of claims 1 to 27 in a unit dosage form.
- 39. A method of treating hyperinsulinemia in a mammal comprising administering to the mammal a therapeutically effective amount of a compound according to any one of claims 1 to 27 in a unit dosage form.
- 40. A method of treating diabetes-induced hypertension in a mammal comprising administering to the mammal a therapeutically effective amount of a compound according to any one of claims 1 to 27 in a unit dosage form.
- 41. A method of treating diabetes-related damage to blood vessels, eyes, kidneys, nerves, autonomic nervous system, skin, connective tissue, or immune system in a mammal comprising administering to the mammal a therapeutically effective amount of a compound according to any one of claims 1 to 27 in a unit dosage form.
- 42. A method of treating obesity in a mammal comprising administering to the mammal a therapeutically effective amount of a compound according to any one of claims 1 to 27 in a unit dosage form.
- 43. A compound according to any one of claims 1 to 27 which is a nicotinic acid derivative.

44. A kit comprising the composition of any one of claims 1 to 27 and instructions for its use in the treatment of a cardiovascular disease, a disease that arises from a thrombotic or prothombotic state in which the coagulation cascade is activated, diabetis, or related diseases.